

WHAT IS CLAIMED IS:

1 1. A method for use in the diagnosis of endometriosis in a subject
2 comprising the steps of:

3 detecting a test amount of a prothymosin gene product in a
4 sample from the subject; and

5 comparing the test amount with a normal amount of the
6 prothymosin gene product in a control sample,

7 whereby a test amount above the normal amount provides a
8 positive indication in the diagnosis of endometriosis.

1 2. The method of claim 1 wherein the sample comprises ectopic
2 endometrial tissue, eutopic endometrial tissue, peritoneal fluid, blood, vaginal
3 secretion or urine.

1 3. The method of claim 1 wherein the prothymosin gene product is
2 prothymosin mRNA or cDNA.

1 4. The method of claim 3 wherein the step of detecting comprises
2 the steps of:

3 contacting the prothymosin mRNA or cDNA with a
4 polynucleotide of at least 7 to about 50 nucleotides in length that specifically
5 hybridizes to the prothymosin mRNA or cDNA and

6 detecting hybridization between the polynucleotide and the
7 mRNA or cDNA.

1 5. The method of claim 4 wherein the polynucleotide comprises
2 DNA or RNA.

1 6. The method of claim 4 wherein the polynucleotide comprises
2 a nucleotide analog selected from the group consisting of phosphorothioates,
3 phosphoramidates, methyl phosphonates, chiral-methyl phosphonates, 2-O-methyl
4 ribonucleotides, and peptide-nucleic acids.

1 7. The method of claim 4 wherein the polynucleotide comprises a
2 detectable moiety, and the step of detecting hybridization comprises detecting the
3 moiety.

1 8. The method of claim 4 wherein the polynucleotide is a primer
2 and the step of detecting hybridization comprises:
3 initiating reverse transcription of prothymosin mRNA with
4 the primer, and
5 detecting a prothymosin mRNA reverse transcript;
6 whereby detection of the reverse transcript indicates that the
7 polynucleotide has specifically hybridized to prothymosin mRNA.

1 9. The method of claim 4 wherein the prothymosin mRNA or
2 cDNA is immobilized and the step of contacting comprises contacting the
3 immobilized mRNA or cDNA with the polynucleotide.

1 10. The method of claim 4 wherein the polynucleotide is
2 immobilized and the step of contacting comprises contacting the immobilized
3 polynucleotide with the prothymosin mRNA or cDNA.

1 11. The method of claim 7 wherein the detectable moiety is a
2 fluorescent label, a radioactive label, an enzymatic label, a biotinyl group, or an
3 epitope recognized by a secondary reporter.

1 12. The method of claim 9 wherein the biological sample is a
2 fixed tissue sample and the step of contacting comprises contacting the
3 polynucleotide with the mRNA or cDNA *in situ* on the fixed tissue sample.

1 ~~13. The method of claim 12 wherein the immobilized~~
2 ~~polynucleotide is comprised within a polynucleotide array.~~

1 14. The method of claim 3 wherein the step of detecting comprises
2 the steps of:

3 amplifying the prothymosin mRNA or cDNA to produce an
4 amplification product and
5 detecting the amplification product.

1 15. The method of claim 14 wherein the step of detecting the
2 amplification product comprises:

3 contacting the amplification product with a polynucleotide of
4 at least 7 to about 50 nucleotides in length that specifically hybridizes to the
5 amplification product, and

6 detecting hybridization between the polynucleotide and the
7 amplification product.
8

1 16. The method of claim 14 wherein the step of detecting the
2 amplification product comprises determining the nucleotide sequence of the
3 amplification product.

1 17. The method of claim 14 wherein the step of detecting the
2 amplification product comprises determining the mass of the amplification product.

1 18. The method of claim 15 wherein the polynucleotide
2 comprises DNA or RNA.

1 19. The method of claim 15 wherein the polynucleotide
2 comprises a nucleotide analog selected from the group consisting of
3 phosphorothioates, phosphoramidates, methyl phosphonates, chiral-methyl
4 phosphonates, 2-O-methyl ribonucleotides, and peptide-nucleic acids.

1 20. The method of claim 15 wherein the polynucleotide comprises
2 a detectable moiety, and the step of detecting hybridization comprises detecting the
3 moiety.

1 21. The method of claim 20 wherein the detectable moiety is a
2 fluorescent label, a radioactive label, an enzymatic label, a biotinyl group, or an
3 epitope recognized by a secondary reporter.

1 22. The method of claim 1 wherein the prothymosin gene product
2 is prothymosin polypeptide.

1 23. The method of claim 22 wherein the step of detecting
2 comprises detecting prothymosin polypeptide by immunoassay.

1 24. The method of claim 22 wherein the step of detecting
2 comprises contacting the sample with an affinity agent that binds to prothymosin
3 polypeptide and detecting binding between the affinity agent and the prothymosin
4 polypeptide.

1 25. The method of claim 22 wherein the step of detecting
2 comprises detecting an analyte in the sample having the mass of prothymosin
3 polypeptide.

1 26. The method of claim 23 wherein the immunoassay is non-
2 competitive immunoassay.

1 27. The method of claim 23 wherein the immunoassay is
2 competitive immunoassay.

1 28. The method of claim 23 wherein the immunoassay comprises
2 detecting binding between the prothymosin polypeptide and an antibody comprising
3 a detectable moiety selected from the group consisting of a fluorescent label, a
4 radioactive label, an enzymatic label, a biotinyl group, and an epitope recognized
5 by a secondary reporter.

1 29. The method of claim 24 wherein the step of detecting binding
2 comprises detecting bound prothymosin polypeptide by mass spectrometry.

1 30. The method of claim 26 wherein the non-competitive
2 immunoassay comprises the steps of:
3 capturing the prothymosin polypeptide from the sample on a
4 solid phase with a first antibody specific for prothymosin polypeptide; and
5 detecting capture of the prothymosin polypeptide by
6 contacting the solid phase with a second antibody specific for prothymosin
7 polypeptide and detecting binding between the second antibody and prothymosin
8 polypeptide.

1 31. The method of claim 26 wherein the non-competitive
2 immunoassay comprises the steps of:
3 binding the prothymosin polypeptide from the sample to a
4 solid phase; and
5 detecting the prothymosin polypeptide by contacting the solid
6 phase with an antibody specific for prothymosin polypeptide and detecting binding
7 between the antibody and prothymosin polypeptide.

1 32. A method for use in the monitoring the progress of
2 endometriosis in a subject comprising the steps of:
3 detecting a first test amount of a prothymosin gene product
4 in a sample from the subject at a first time;
5 detecting a second test amount of the prothymosin gene
6 product in a sample from the subject at a second, later time; and
7 comparing the first test amount with the second test amount,
8 whereby an increase in the amount between the first time and
9 the second time indicates progression of endometriosis and a decrease in the
10 amount between the first time and the second time indicates remission of
11 endometriosis.

1 33. A kit comprising a compound that binds a prothymosin gene
2 product and instructions to (1) use the compound for detecting prothymosin in a
3 patient sample, and (2) to diagnose endometriosis based on an elevated amount of
4 the prothymosin gene product in the sample compared with a normal amount of
5 prothymosin.

1 34. The kit of claim 33 wherein the prothymosin gene product is
2 prothymosin mRNA or cDNA and the compound is a polynucleotide that
3 hybridizes with prothymosin mRNA or cDNA under stringent conditions.

1 35. The kit of claim 33 wherein the prothymosin gene product is
2 prothymosin polypeptide and the compound is an antibody that specifically binds to
3 prothymosin polypeptide.

1 36. A method for use in the diagnosis of endometriosis in a subject
2 comprising detecting a prothymosin gene product in endometriotic tissue from the
3 subject *in vivo*, whereby detection of the gene product provides a positive
4 indication in the diagnosis of endometriosis.

1 37. The method of claim 36 comprising administering to the
2 subject a compound that specifically binds to a prothymosin gene product and
3 detecting binding between the compound and the prothymosin gene product.

1 38. The method of claim 37 wherein the compound comprises a
2 gamma-emitting or positron-emitting radioisotope and binding is detected by
3 detecting the radioisotope by camera imaging or Geiger counter.

1 39. The method of claim 37 wherein the compound comprises a
2 paramagnetic isotope and binding is detected by detecting the paramagnetic isotope
3 by magnetic resonance imaging ("MRI").

1 40. The method of claim 37 wherein the compound is a
2 polynucleotide that specifically hybridizes to prothymosin mRNA.

1 41. The method of claim 37 wherein the compound is an
2 antibody that specifically hybridizes to prothymosin polypeptide.

1 42. A method for the treatment of endometriosis in a subject
2 comprising:
3 administering to the subject a probe comprising a detectable
4 label and a ligand that specifically binds a prothymosin gene product, to allow
5 binding between the probe and the prothymosin gene product;
6 identifying an endometriotic lesion *in situ* by locating bound
7 label; and
8 excising the endometriotic lesion.

1 43. The method of claim 42 comprising:
2 administering the probe into the peritoneum of the subject,
3 wherein the probe comprises an antibody ligand that specifically binds prothymosin
4 and a radioactive label;
5 identifying an endometriotic lesion *in situ* by locating bound
6 probe with a Geiger counter; and
7 excising the endometriotic lesion laparoscopically.

1 44. A screening method for determining whether a compound
2 modulates the expression of a prothymosin gene product in an endometrial cell
3 comprising the steps of:
4 contacting the cell with the compound; and
5 determining whether expression of the prothymosin gene
6 product is different that expression in a control cell which has not been contacted
7 with the compound;
8 whereby a difference between expression in the endometrial
9 cell and the control cell indicates that the agent modulates expression of the
10 prothymosin gene product.

1 45. The method of claim 44 wherein:
2 the endometrial cell is comprised within endometriotic tissue
3 cultured as a xenograft in a mouse;
4 the step of contacting comprises administering the compound
5 to the mouse;
6 the step of determining comprises *in vitro* determination of
7 expression of the gene product after removing the tissue from the mouse.

1 46. A method for the treatment of endometriosis in a subject
2 comprising the step of administering to the subject a compound that decreases
3 prothymosin activity in eutopic endometrial tissue or ectopic endometrial tissue in
4 the subject.

1 47. The method of claim 46 wherein the compound inhibits
2 expression of prothymosin mRNA.

1 48. The method of claim 46 wherein the compound inhibits
2 activity of prothymosin protein.

1 49. The method of claim 46 wherein the compound is a small
2 organic molecule.

1 50. The method of claim 46 wherein the compound is
2 administered intraperitoneally.

1 51. The method of claim 47 wherein the compound comprises an
2 inhibitory polynucleotide comprising a sequence of at least 7 nucleotides identical
3 or complementary to prothymosin mRNA sequence, wherein the inhibitory
4 polynucleotide inhibits transcription, processing or translation of prothymosin
5 mRNA.

1 52. The method of claim 51 wherein the inhibitory
2 polynucleotide is a polynucleotide comprising an antisense sequence of at least 7
3 nucleotides that specifically hybridizes to a nucleotide sequence within
4 prothymosin mRNA, whereby the polynucleotide inhibits the activity of the
5 prothymosin mRNA.

1 53. The method of claim 51 wherein the inhibitory
2 polynucleotide is a ribozyme that cleaves prothymosin mRNA.

1 54. The method of claim 52 wherein the antisense sequence is
2 between 10 and 50 nucleotides in length.

1 55. The method of claim 52 wherein the polynucleotide
2 comprises a nucleotide analog selected from phosphorothioates, phosphoramidates,
3 methyl phosphonates, chiral-methyl phosphonates, 2-O-methyl ribonucleotides and
4 peptide-nucleic acids.

1 56. The method of claim 52 wherein the step of providing the
2 cells with the polynucleotide comprises transfecting the cells with an expression
3 vector comprising expression control sequences operatively linked to a nucleotide
4 sequence encoding the antisense polynucleotide, whereby the vector expresses the
5 polynucleotide.